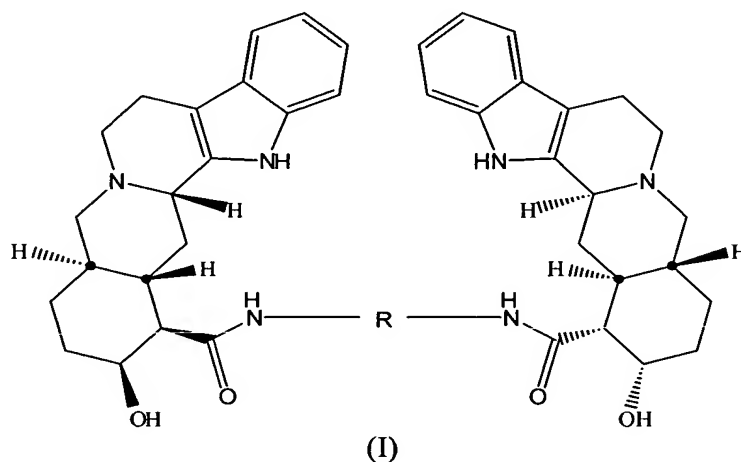


**What is claimed is:**

1. A method of treating or preventing an  $\alpha_2$  adrenergic receptor mediated condition or disorder comprising:  
providing a compound according to formula (I)



wherein R is a linker molecule that affords activity of the compound as an  $\alpha_2$  adrenergic receptor antagonist; and

- administering to a patient an effective amount of the compound to treat or prevent the  $\alpha_2$  adrenergic receptor mediated condition or disorder.

2. The method according to claim 1, wherein the  $\alpha_2$  adrenergic receptor mediated condition or disorder is an  $\alpha_{2a}$  adrenergic receptor mediated condition or disorder.

3. The method according to claim 2, wherein the  $\alpha_{2a}$  adrenergic receptor mediated condition or disorder is selected from the group consisting of hypertension, hypotension, erectile dysfunction, pain, glaucoma, alcohol and drug withdrawal, rheumatoid arthritis, ischemia, migraine, cognitive deficiency, spasticity, diarrhea, and nasal congestion.

4. The compound according to claim 2, wherein the compound exhibits selectivity in binding an  $\alpha_{2a}$  adrenergic receptor over an  $\alpha_{2b}$  adrenergic receptor.

5. The method according to claim 1, wherein the  $\alpha_2$  adrenergic receptor mediated condition or disorder is an  $\alpha_{2c}$  adrenergic receptor mediated condition or disorder.

5 6. The method according to claim 5, wherein the  $\alpha_{2c}$  adrenergic receptor mediated condition or disorder is Raynaud's disease.

7. The method according to claim 5, wherein the compound exhibits selectivity in binding an  $\alpha_{2c}$  adrenergic receptor over an  $\alpha_{2b}$  adrenergic receptor.  
10

8. The method according to claim 1, wherein R has a length of about 2.5 Å to about 45 Å.

15 9. The method according to claim 8 wherein R has a length of about 2.5 Å to about 5 Å.

10. The method according to claim 8 wherein R has a length of about 23 Å to about 29 Å.

20 11. The method according to claim 1, wherein R is either:  
(i) a straight or branched chain alkyl, alkenyl, alkynyl comprising at least 2 carbon atoms in a main chain thereof, or  
(ii) a straight or branched chain alkyl, alkenyl, alkynyl  
25 comprising at least 2 carbon atoms in a main chain thereof and an X group within the main chain and/or a Y group as a substituent linked to a carbon atom in the main chain,

with X being —O—, carbonyl, —NR<sup>1</sup>— with R<sup>1</sup> being H or an alkyl, —C(O)NHR<sup>1</sup>— with R<sup>1</sup> being an alkyl, —S—, sulfoxide, sulfonyl, or a cyclic or multicyclic ring with or without hetero atoms as ring members and including, optionally, one or more substitutions on the ring structure, and  
30

with Y being —OH, —NO<sub>2</sub>, —CN, —C(O)H, —SH, a primary, secondary, or tertiary amine, a carboxylic acid, an ester, a keto group, —SO<sub>2</sub>NH<sub>2</sub>, or —SO<sub>2</sub>NHR<sup>2</sup> with R<sup>2</sup> being an alkyl, or

(iii) a cyclic or multicyclic ring with or without hetero atoms as ring members and including, optionally, one or more substitutions on the ring structure(s).

12. The method according to claim 11, wherein R is a straight chain alkyl.

13. The method according to claim 12, wherein R is a straight chain C2 to C36 alkyl.

14. The method according to claim 12, wherein R is a straight chain C3 to C24 alkyl.

15. The method according to claim 11, wherein R is a straight chain alkyl comprising at least 4 carbon atoms and an X group within the straight chain, wherein X is a —O—.

16. The method according to claim 15, wherein R is —(CH<sub>2</sub>CH<sub>2</sub>-O-CH<sub>2</sub>CH<sub>2</sub>)<sub>n</sub>— wherein n is an integer from 1 to 6.

17. The method according to claim 15, wherein R is —(CH<sub>2</sub>CH<sub>2</sub>-O-CH<sub>2</sub>CH<sub>2</sub>-O-CH<sub>2</sub>CH<sub>2</sub>)<sub>n</sub>— wherein n is an integer from 1 to 4.

18. The method according to claim 11, wherein R is a straight chain alkyl comprising at least 5 carbon atoms and an X group within the straight chain, wherein X is —C(O)NHR<sup>1</sup>— with R<sup>1</sup> being an alkyl.

19. The method according to claim 11, wherein R is —(CH<sub>2</sub>-NHC(O))<sub>n</sub>-CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>-(C(O)NH-CH<sub>2</sub>)<sub>n</sub>— wherein each n is independently an integer from 1 to 3.

20. The method according to claim 11, wherein R is a straight chain alkenyl comprising at least 5 carbon atoms in a main chain thereof and an X group within the main chain, wherein X is  $\text{—C(O)NHR}^1\text{—}$  with  $\text{R}^1$  being an alkyl.

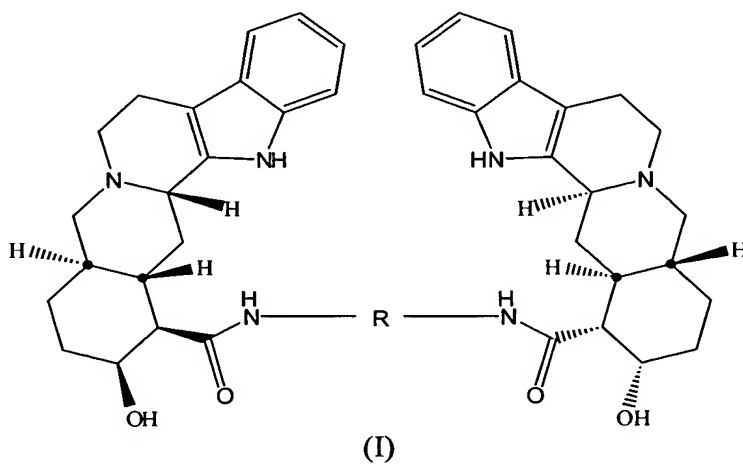
21. The method according to claim 20, wherein R is a cis isomer or a trans isomer of  $\text{—(CH}_2\text{—NHC(O))}_n\text{—CH}_2\text{—CH=CH—CH}_2\text{—(C(O)NH—CH}_2\text{)}_n\text{—}$  wherein each n is independently an integer from 1 to 3.

22. The method according to claim 21, wherein R is a cis isomer.

23. The method according to claim 21, wherein R is a trans isomer.

24. A method of modulating the activity of an  $\alpha_{2a}$  adrenergic receptor comprising:

providing a compound according to formula (I)



wherein R is a linker molecule that affords activity of the compound as an  $\alpha_{2a}$  adrenergic receptor antagonist; and

contacting an  $\alpha_{2a}$  adrenergic receptor with the compound under conditions effective to modulate the activity of the  $\alpha_{2a}$  adrenergic receptor.

25. The method according to claim 24, wherein the compound exhibits selectivity in binding an  $\alpha_{2a}$  adrenergic receptor over an  $\alpha_{2b}$  adrenergic receptor.

5                    26. The method according to claim 24, wherein R has a length of about 2.5 Å to about 45 Å.

27. The method according to claim 24, wherein R is either:

(i) a straight or branched chain alkyl, alkenyl, alkynyl  
10 comprising at least 2 carbon atoms in a main chain thereof, or

(ii) a straight or branched chain alkyl, alkenyl, alkynyl comprising at least 2 carbon atoms in a main chain thereof and an X group within the main chain and/or a Y group as a substituent linked to a carbon atom in the main chain,

15                    with X being —O—, carbonyl, —NR<sup>1</sup>— with R<sup>1</sup> being H or an alkyl, —C(O)NHR<sup>1</sup>— with R<sup>1</sup> being an alkyl, —S—, sulfoxide, sulfonyl, or a cyclic or multicyclic ring with or without hetero atoms as ring members and including, optionally, one or more substitutions on the ring structure, and

20                    with Y being —OH, —NO<sub>2</sub>, —CN, —C(O)H, —SH, a primary, secondary, or tertiary amine, a carboxylic acid, an ester, a keto group, —SO<sub>2</sub>NH<sub>2</sub>, or —SO<sub>2</sub>NHR<sup>2</sup> with R<sup>2</sup> being an alkyl, or

(iii) a cyclic or multicyclic ring with or without hetero atoms as ring members and including, optionally, one or more substitutions on the  
25 ring structure(s).

28. The method according to claim 27, wherein R is a straight chain alkyl.

30                    29. The method according to claim 27, wherein R is a straight chain alkyl comprising at least 4 carbon atoms and an X group within the straight chain, wherein X is a —O—.

30. The method according to claim 27, wherein R is a straight chain alkyl comprising at least 5 carbon atoms and an X group within the straight chain, wherein X is  $\text{—C(O)NHR}^1\text{—}$  with  $\text{R}^1$  being an alkyl.

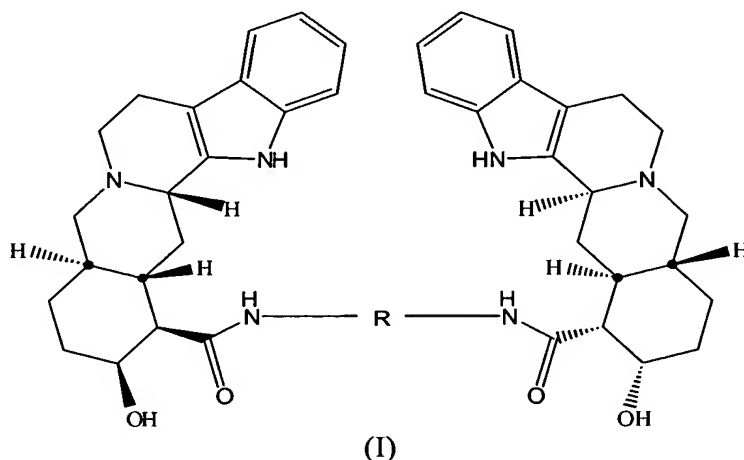
31. The method according to claim 27, wherein R is  $\text{—(CH}_2\text{—NHC(O))}_n\text{—CH}_2\text{—CH}_2\text{—CH}_2\text{—CH}_2\text{—(C(O)NH—CH}_2\text{)}_n\text{—}$  wherein each n is independently an integer from 1 to 3.

32. The method according to claim 27, wherein R is a straight chain alkenyl comprising at least 5 carbon atoms in a main chain thereof and an X group within the main chain, wherein X is  $\text{—C(O)NHR}^1\text{—}$  with  $\text{R}^1$  being an alkyl.

33. The method according to claim 32, wherein R is a cis isomer or a trans isomer of  $\text{—(CH}_2\text{—NHC(O))}_n\text{—CH}_2\text{—CH=CH—CH}_2\text{—(C(O)NH—CH}_2\text{)}_n\text{—}$  wherein each n is independently an integer from 1 to 3.

34. A method of modulating the activity of an  $\alpha_{2c}$  adrenergic receptor comprising:

providing a compound according to formula (I)



wherein R is a linker molecule that affords activity of the compound as an  $\alpha_{2c}$  adrenergic receptor antagonist; and

contacting an  $\alpha_{2c}$  adrenergic receptor with the compound under conditions effective to modulate the activity of the  $\alpha_{2c}$  adrenergic receptor.

35. The compound according to claim 34, wherein the yohimbine dimer exhibits selectivity in binding an  $\alpha_{2c}$  adrenergic receptor over an  $\alpha_{2b}$  adrenergic receptor.

5 36. The method according to claim 34, wherein the compound exhibits selectivity in binding an  $\alpha_{2c}$  adrenergic receptor over an  $\alpha_{2a}$  adrenergic receptor.

10 37. The method according to claim 34, wherein R has a length of about 2.5 Å to about 45 Å.

38. The method according to claim 34, wherein R is either:

(i) a straight or branched chain alkyl, alkenyl, alkynyl comprising at least 2 carbon atoms in a main chain thereof, or

15 (ii) a straight or branched chain alkyl, alkenyl, alkynyl comprising at least 2 carbon atoms in a main chain thereof and an X group within the main chain and/or a Y group as a substituent linked to a carbon atom in the main chain,

20 with X being —O—, carbonyl, —NR<sup>1</sup>— with R<sup>1</sup> being H or an alkyl, —C(O)NHR<sup>1</sup>— with R<sup>1</sup> being an alkyl, —S—, sulfoxide, sulfonyl, or a cyclic or multicyclic ring with or without hetero atoms as ring members and including, optionally, one or more substitutions on the ring structure, and

25 with Y being —OH, —NO<sub>2</sub>, —CN, —C(O)H, —SH, a primary, secondary, or tertiary amine, a carboxylic acid, an ester, a keto group, —SO<sub>2</sub>NH<sub>2</sub>, or —SO<sub>2</sub>NHR<sup>2</sup> with R<sup>2</sup> being an alkyl, or

(iii) a cyclic or multicyclic ring with or without hetero atoms as ring members and including, optionally, one or more substitutions on the ring structure(s).

30 39. The method according to claim 38, wherein R is a straight chain alkyl.

40. The method according to claim 38, wherein R is a straight chain alkyl comprising at least 4 carbon atoms and an X group within the straight chain, wherein X is a —O—.

5 41. The method according to claim 38, wherein R is a straight chain alkyl comprising at least 5 carbon atoms and an X group within the straight chain, wherein X is —C(O)NHR<sup>1</sup>— with R<sup>1</sup> being an alkyl.

42. The method according to claim 38, wherein R is  
10 —(CH<sub>2</sub>-NHC(O))<sub>n</sub>-CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>-CH<sub>2</sub>-(C(O)NH-CH<sub>2</sub>)<sub>n</sub>— wherein each n is independently an integer from 1 to 3.

43. The method according to claim 38, wherein R is a straight chain alkenyl comprising at least 5 carbon atoms in a main chain thereof and an X  
15 group within the main chain, wherein X is —C(O)NHR<sup>1</sup>— with R<sup>1</sup> being an alkyl.

44. The method according to claim 43, wherein R is a cis isomer or a trans isomer of  
—(CH<sub>2</sub>-NHC(O))<sub>n</sub>-CH<sub>2</sub>-CH=CH-CH<sub>2</sub>-(C(O)NH-CH<sub>2</sub>)<sub>n</sub>— wherein each n is  
20 independently an integer from 1 to 3.